SEQUENCE LISTING

(1) GENERAL INFORMATION:

(i) APPLICANT: IMMER, Hansueli

FOR\$SMANN, Wolf-Georg

ADERMANN, Knut KLESSEN, Christian



(ii) TITLE OF INVENTION: PROCESS AND INTERMEDIATE PRODUCTS FOR PREPARING CARDIODILATIN FRAGMENTS AND HIGHLY PURIFIED CARDIODILATIN FRAGMENTS

- (iii) NUMBER OF SEQUENCES: 11
- (iv) CORRESPONDENCE ADDRESS:
 - (A) ADDRESSEE: Nikaido, Marmelstein, Murray & Oram
 - (B) STREET: 655 Fitteenth Street, N.W., Suite 330
 - (C) CITY: Washington
 - (D) STATE: D.C.
 - (E) COUNTRY: USA
 - (F) ZIP: 20005-4810
 - (v) COMPUTER READABLE FORM:

 - (A) MEDIUM TYPE: Floopy disk
 (B) COMPUTER: IBM PC compatible
 (C) OPERATING SYSTEM: PC-DOS/MS-DOS
 - (D) SOFTWARE: PatentIn Release #1.0, Version #1.30
- (vi) CURRENT APPLICATIØN DATA:
 - (A) APPLICATION/NUMBER:\ US 09/027,777
 - (B) FILING DATZ: 23-FEB-1998
 - (C) CLASSIFICATION:
- (vii) PRIOR APPLICATION DATA:
 - (A) APPLICATION NUMBER: DE 44 20 381.0
 - (B) FILING DATE: 02-JUN-1994
- vii) PRIOR APPLICATION DATA:
 - (A) APPLICATION NUMBER: DE\195 13 784.1
 - (B) FILING DATE: 10-APR-1995
- (viii) ATXORNEY/AGENT INFORMATION:
 - (A) NAME: Kitts, Monica C.
 - (B) REGISTRATION NUMBER: 36, 105
 - (C) REFERENCE/DOCKET NUMBER: P1614-8019
 - TELECOMMUNICATION INFORMATION:
 - (A) TELEPHONE: (202) 638-5000
 - (B) TELEFAX: (202) 368-4810
- (2) INFORMATION FOR SEQ ID NO:1:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 17 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

Cys Phe Gly Gly Arg Met Asp Arg Ile Gly Ala Gln Ser Gly Leu Gly
1 5 10 15

Cys

- (2) INFORMATION FOR SEQ ID NO:2:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 15 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

Leu Arg Ala Leu Leu Thr Ala Pro Arg Ser Leu Arg Arg Ser Ser 1 10 15

- (2) INFORMATION FOR SEQ ID NO:3:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 5 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

Asn Ser Phe Arg Tyr 5

- (2) INFORMATION FOR SEQ ID NO:4:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 37 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear

(1)

- (ii) MOLECULE TYPE: peptide
- (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION: 1..15
- (D) OTHER INFORMATION: /note= "Xaa chain is an amino acid chain of 0 15 amino acids."
 - (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION: 33..37
- (D) OTHER INFORMATION: /note= "Xaa chain is an amino acid chain of 0 5 amino acids."
 - (ix) FEATURE:
 - (A) NAME/KEY: Cross-links
 - (B) LOCATION: 16..32
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

Phe Gly Gly Arg Met Asp Arg Ile Gly Ala Gln Ser Gly Leu Gly Cys 20 25 30

Xaa Xaa Xaa Xaa 35

- (2) INFORMATION FOR SEQ ID NO:5:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 32 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY: Cross-links
 - (B) LOCATION: 11..27
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:

Thr Ala Pro Arg Ser Leu Arg Arg Ser Ser Cys Phe Gly Gly Arg Met
1 5 10 15

Asp Arg Ile Gly Ala Gln Ser Gly Leu Gly Cys Asn Ser Phe Arg Tyr 20 25 30

- (2) INFORMATION FOR SEQ ID NO:6:
 - (i) SEQUENCE CHARACTERISTICS:

4

- (A) LENGTH: 14 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 1
- (D) OTHER INFORMATION: /product= "Modified by a Boc protecting group and a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 4
- (D) OTHER INFORMATION: /note= "Modified by a tBu protecting group and a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 7
- (D) OTHER INFORMATION: /note= "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 8
- (D) OTHER INFORMATION: /note= "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 9
- (D) OTHER INFORMATION: /note = "Modified by a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 10
- (D) OTHER INFORMATION: /note= "Modified by a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 11
- (D) OTHER INFORMATION: /note = "Modified by a Acm protecting group"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

Thr Ala Pro Arg Ser Leu Arg Arg Ser Ser Cys Phe Gly Gly 1

- (2) INFORMATION FOR SEQ ID NO:7:
 - (i) SEQUENCE CHARACTERISTICS:



- (A) LENGTH: 12 amino acids
- (B) TYPE: amino acid
- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 1
- (D) OTHER INFORMATION: /product= "Modified by a Fmoc protecting group and a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 3
- (D) OTHER INFORMATION: /product= "Modified by a OtBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 4
- (D) OTHER INFORMATION: /product = "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 8
- (D) OTHER INFORMATION: $\product=$ "Modified by a Trt protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 9
- (D) OTHER INFORMATION: /product = "Modified by a tBu protecting group"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:

Arg Met Asp Arg Ile Gly Ala Gln Ser Gly Leu Gly 1 5 10

- (2) INFORMATION FOR SEQ ID NO:8:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 6 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 1
 - (D) OTHER INFORMATION: /note= "Modified by an Fmoc protecting

group or H"

- (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 1
- (D) OTHER INFORMATION: /product= "Modified by a Trt protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 2
- (D) OTHER INFORMATION: /product = "Modified by a Trt protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 3
- (D) OTHER INFORMATION: /product= "Modified by a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 5
- (D) OTHER INFORMATION: /product = "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 6
- (D) OTHER INFORMATION: /product = "Modified by a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 6
- (D) OTHER INFORMATION: /note= "Modified by an O-OH or an O-OtBu protecting group"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

Cys Asn Ser Phe Arg Tyr 1 5

- (2) INFORMATION FOR SEQ ID NO:9:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 18 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 1
 - (D) OTHER INFORMATION: /note= "Modified by an H or an Fmoc

9

protecting group"

- (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 1
- (D) OTHER INFORMATION: /product= "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 3
- (D) OTHER INFORMATION: /product= "Modified by a OtBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 4
- (D) OTHER INFORMATION: /product= "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 8
- (D) OTHER INFORMATION: /product= "Modified by a Trt protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 9
- (D) OTHER INFORMATION: /product = "Modified by a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 13
- (D) OTHER INFORMATION: /product= "Modified by a Trt protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 14
- (D) OTHER INFORMATION: /product= "Modified by a Trt protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 15
- (D) OTHER INFORMATION: /product= "Modified by a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 17
- (D) OTHER INFORMATION: /product= "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 18



- (D) OTHER INFORMATION: /product = "Modified by a tBu protecting group and an OtBu protecting group"
 - (xi) SEQUENCE DESCRIPTION: SEO ID NO:9:

Arg Met Asp Arg Ile Gly Ala Gln Ser Gly Leu Gly Cys Asn Ser Phe 1 5 10 15

Arg Tyr

- (2) INFORMATION FOR SEQ ID NO:10:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 32 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 1
- (D) OTHER INFORMATION: /product= "Modified by a Boc protecting group and a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 4
- (D) OTHER INFORMATION: /product = "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 5
- (D) OTHER INFORMATION: /product = "Modified by a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 7
- (D) OTHER INFORMATION: /product = "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 8
- (D) OTHER INFORMATION: /product= "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 9
- (D) OTHER INFORMATION: /product = "Modified by a Bu protecting group"



- (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 10
- (D) OTHER INFORMATION: /product= "Modified by a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 11
- (D) OTHER INFORMATION: /product = "Modified by a Acm protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 15
- (D) OTHER INFORMATION: /product= "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 17
- (D) OTHER INFORMATION: /product= "Modified by a OtBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 18
- (D) OTHER INFORMATION: /product = "Modified by a Pbf protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 22
- (D) OTHER INFORMATION: /product = "Modified by a Trt protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 23
- (D) OTHER INFORMATION: /product= "Modified by a tBu protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 27
- (D) OTHER INFORMATION: /product = "Modified by a Trt protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 28
- (D) OTHER INFORMATION: /product = "Modified by a Trt protecting group"
 - (ix) FEATURE:
 - (A) NAME/KEY: Modified-site
 - (B) LOCATION: 29
- (D) OTHER INFORMATION: /product= "Modified by a tBu protecting group"





(ix) FEATURE:

- (A) NAME/KEY: Modified-site
- (B) LOCATION: 31
- (D) OTHER INFORMATION: /product = "Modified by a Pbf protecting group"

(ix) FEATURE:

- (A) NAME/KEY: Modified-site
- (B) LOCATION: 32
- (D) OTHER INFORMATION: /product= "Modified by a tBu protecting group and an OtBu protecting group"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:10:

Thr Ala Pro Arg Ser Leu Arg Arg Ser Ser Cys Phe Gly Gly Arg Met
1 5 10 15

Asp Arg Ile Gly Ala Gln Ser Gly Leu Gly Cys Asn Ser Phe Arg Tyr 20 25 30

(2) INFORMATION FOR SEQ ID NO:11:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 32 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide

(ix) FEATURE:

- (A) NAME/KEY: Modified-site
- (B) LOCATION: 11
- (D) OTHER INFORMATION: /product = "Modified by a Acm protecting group"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:11:

Thr Ala Pro Arg Ser Leu Arg Arg Ser Ser Cys Phe Gly Gly Arg Met $1 \hspace{1.5cm} 5 \hspace{1.5cm} 10 \hspace{1.5cm} 15$

Asp Arg Ile Gly Ala Gln Ser Gly Leu Gly Cys Asn Ser Phe Arg Thr 20 25 30

D